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Title: FLAVANONE COMPOUND AND USES THEREOF
Inventor(s): KUMAZAWA

Docket No.: 50266-00001

David F. Dockery (303) 338-0997

Fig. 1

Compound 1 - Physicochemical Properties

| Appearance | white powder |
|---|--|
| Molecular formula | C ₂₅ H ₂₈ O ₆ |
| ESI-MS (m/z) | |
| Positive: | 425.3 (M+H) ⁺ |
| Negative: | 423.5 (M-H) |
| UV $\lambda_{\max}^{\text{MeOH}}$ nm (ϵ) | 288.5 (19,674) |
| IR (KBr) cm ⁻¹ | 3420, 2960, 2920, 1680, 1600, 1160 |
| $[\alpha]_0^{24}$ (c 0.2, MeOH) | ~5.8° |
| $[\alpha]_{D}^{25}$ (c 0.2, CHCl ₃) | +20.0° |
| MP | 80∼83°C |
| CD (MeOH) θ /deg | 8470 (330), -46997 (290), 12833 (255) |

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Fig. 2

NMR Data (Compound 1 in Acetone-ds)

| Position | ¹³ C | | | ¹H |
|----------|-----------------|-----------------|-------|-----------------------------|
| 2 | 77.21 | СН | 5.62 | (1H, dd), J=2.9, 11.8 Hz |
| 3 | 43.26 | CH ₂ | 2.66 | (1H, dd), $J=2.9$, 17.2 Hz |
| • | | | 3.17 | (1H, dd), J=11.8, 17.2 Hz |
| 4 | 197.59 | С | | |
| 5 | 165.29 | С | 12.19 | (-OH, s) |
| . 6 | 95.83 | CH: | 5.96 | (1H, s) |
| -7 | 167.27 | С | | |
| 8 | 96.79 | СН | 5.96 | (1H, s) |
| 9 | 164.65 | С | | |
| 10 | 103.12 | С | , | |
| 1' | 129.77 | С | | |
| 2' | 127,71 | С | | |
| 3, | 144.08 | С | | |
| 4' | 145.56 | С | | |
| 5' | 113.49 | СН | 6.82 | (1H, d), J=8.3 Hz |
| 6' | 118.55 | СН | 6.96 | (1H, d), √=8.3 Hz |
| 1" | 25.15 | CH ₂ | 3.55 | (2H, d), J=6.3 Hz |
| 2" | 124.15 | СН | 5.18 | (1H, t), J=6.3 Hz |
| 3" | 135.33 | С | | <u> </u> |
| 4'' | 16.34 | СН₃ | 1.68 | (3H, s) |
| 5" | 40.37 | CH ₂ | 1.97 | (2H, t), J=6.2 Hz |
| 6'' | 27.34 | CH ₂ | 2.05 | (2H, m) |
| 7" | 125.02 | СН | 5.06 | (1H, dt), J=1.9, 6.2 Hz |
| 8" | 131.74 | С | | |
| 9" | 25.75 | CH₃ | 1.60 | (3H, s) |
| 10" | 17.68 | CH ₃ | 1.55 | (3H, s) |

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Fig. 3

Compound 2 - Physicochemical Properties

| | · |
|---------------------------------------|---------------------------------------|
| Appearance | yellow powder |
| Molecular formula | $C_{25H_{28}O_{6}}$ |
| ESI-MS (m/z) | |
| Positive: | 425.0 (M+H) ⁺ |
| Negative: | 423.3 (M-H) |
| HRFAB-MS (m/z) | |
| calcd.: | 425.1965(M+H) ⁺ |
| found: | 425.1968(M+H) ⁺ |
| UV λ max nm (ε) | 288.0 (17,935) |
| IR (KBr) cm ⁻¹ | 3360, 2960, 2920, 1680, 1600 |
| $[\alpha]_{\rm D}^{25}$ (c 0.2, MeOH) | -17.8° |
| MP | 123∼126°C |
| CD (MeOH) θ /deg | 11444 (332), -30028 (292), 7139 (254) |

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Fig. 4

NMR Data (Compound 2 in Acetone-de)

| | NMH | Data (C | ompound | 2 in Acetone-d ₆) | |
|----------|-----------------|-----------------|------------------------------------|-----------------------------------|----|
| Position | ¹³ C | | ion ¹³ C ¹ H | | ¹H |
| 2 | 80.13 | СН | 5.35 | (1H, dd), <i>J</i> =2.9, 12.2 Hz | |
| 3 | 43.54 | CH ₂ | 2.74 | (1H, dd), J=2.9, 17.1 Hz | |
| | | | 3.12 | (1H, dd), <i>J</i> =12.2, 17.1 Hz | |
| 4 | 197.19 | Ċ | | | |
| 5 | 165.24 | С | 12.17 | (-OH, s) | |
| 6 | 95.81 | СН | 5.95 | (1H, s) | |
| 7 | 167.28 | С | | · | |
| 8 | 96.74 | СН | 5.95 | (1H, s) | |
| 9 | 164.31 | C | | | |
| 10 | 103.22 | С | | | |
| 1' | 130.58 | С | | | |
| 2' | 112.07 | СН | 6.91 | (1H, d), J=2.2 Hz | |
| . 3' | 145.26 | С | | | |
| 4' | 144.30 | С | | | |
| 5' | 129.02 | С | | | |
| 6, | 119.97 | СН | 6.81 | (1H, d), J=2.2 Hz | |
| 1" | 28.83 | CH ₂ | 3.38 | (2H, d), <i>J</i> =7.3 Hz | |
| 2'' | 123.36 | СН | 5.38 | (1H, m) | |
| 3" | 136.39 | C . | | | |
| 4'' | 16.21 | CH3 | 1.73 | (3H, s) | |
| 5'' | 40.45 | CH ₂ | 2.06 | (2H, t), <i>J</i> =7.5 Hz | |
| 6" | 27.37 | CH ₂ | 2.12 | (2H, td), J=6.8, 7.5 Hz | |
| 7'' | 125.07 | СН | 5.12 | (1H, tq), J=1.5, 6.8 Hz | |
| 8" | 131.70 | С | | | |
| 9" | 25.80 | CH₃ | 1.63 | (3H, s) | |
| 10" | 17.71 | CH ₃ | 1.57 | (3H, s) | |

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Title: FLAVANONE COMPOUND AND USES THEREOF

Inventor(s): KUMAZAWA Docket No.: 50266-00001

David F. Dockery

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Fig. 5

Compound 3 - Physicochemical Properties

| Appearance | yellow powder |
|---|--------------------------------------|
| Molecular formula | $C_{25H_{28}O_{6}}$ |
| ESI-MS (m/z) | |
| Positive: | 425.1 (M+H) ⁺ |
| Negative: | 423.2 (M-H) |
| UV λ_{\max}^{MeOH} nm ($arepsilon$) | 291.5 (16,833) |
| IR (KBr) cm ⁻¹ | 3380, 2960, 2920, 1680, 1600, 1450 |
| $[\alpha]_{0}^{23}$ (c 0.77, MeOH) | -3.94° |
| $[\alpha]_{\rm D}^{25}$ (c 0.2, CHCl ₃) | -6.5° |
| MP | 172∼175°C |
| CD (MeOH) θ /deg | 9773 (335), -26940 (293), 3399 (255) |

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Title: FLAVANONE COMPOUND AND USES THEREOF
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David F. Dockery (303) 338-0997

Fig. 6

| · · | NMR | Data (C | ompound | 3 in Acetone-d ₆) | | |
|----------|-----------------|-----------------|---------|-----------------------------------|--|--|
| Position | ¹³ C | | · | ¹ H | | |
| 2 | 79.85 | СН | 5.35 | (1H, dd), J=3.0, 12.7 Hz | | |
| . 3 | 43.64 | CH ₂ | 2.71 | (1H, dd), J=3.0, 17.1 Hz | | |
| | | | 3.12 | (1H, dd), <i>J</i> =12.7, 17.1 Hz | | |
| 4 | 197.26 | C | | | | |
| 5 | 162.24 | C | 12.46 | (-OH, s) | | |
| 6 | 108.99 | C | | | | |
| 7 | 164.76 | C | | | | |
| 8 | 95.27 | СН | 6.03 | (1H, s) | | |
| 9 | 161.91 | C | | | | |
| .10 | 103.09 | С | | | | |
| 1' | 131.69 | С | | , | | |
| 2' | 114.67 | CH | 7.03 | (1H, s) | | |
| 3' | 145.95 | C | | | | |
| 4' | 146.28 | C | | - | | |
| 5' | 115.97 | СН | 6.86 | (1H, s) | | |
| 6' | 119.18 | СН | 6.86 | (1H, s) | | |
| .1" | 21.53 | CH ₂ | 3.26 | (2H, d), <i>J</i> =7.3 Hz | | |
| 2" | 123.44 | СН | 5.26 | (1H, td), J=1.0, 7.3 Hz | | |
| 3" | 134.96 | С | | | | |
| 4" | 16.18 | CH₃ | 1.76 | (3H, s) | | |
| 5" | 40.46 | CH ₂ | 1.95 | (2H, t), J=7.5 Hz | | |
| 6" | 27.38 | CH₂ | 2.05 | (2H, m) | | |
| 7" | 125.12 | СН | 5.08 | (1H, tt), J=1.0, 5.4 Hz | | |
| 8" | 131.54 | С | | | | |
| 9" | 25.77 | CH3 | 1.62 | (3H, s) | | |
| 10" | 17.66 | CH ₃ | 1.56 | (3H, s) | | |

Serial No.: PCT/JP2004/008964 Page 7 of 8 Title: FLAVANONE COMPOUND AND USES THEREOF

Inventor(s): KUMAZAWA Docket No.: 50266-00001 David F. Dockery

Pavid F. Dockery (303) 338-0997

Fig. 7

Compound 4 - Physicochemical Properties

| Appearance | light brown gum | |
|---|--|--|
| Molecular formula | C ₃₀ H ₃₆ O ₆ | |
| FAB-MS (m/z) | | |
| Positive: | 493.3 (M+H) [†] | |
| UV λ_{\max}^{MeOH} nm (ε) | 292.0 (20,418) | |
| IR (KBr) cm ⁻¹ | 3400, 2960, 2920, 1640, 1600 | |
| $[\alpha]_{\rm D}^{24}$ (c 0.2, MeOH) | +1.8° | |
| $[\alpha]_{\rm D}^{\rm 25}$ (c 0.2, CHCl ₃) | +26.5° | |
| CD (MeOH) θ /deg | 8810 (335), -30708 (293), 10510 (257) | |

Title: FLAVANONE COMPOUND AND USES THEREOF Inventor(s): KUMAZAWA

Docket No.: 50264, 2020.

Docket No.: 50266-00001 David F. Dockery

(303) 338-0997

Fig. 8

NMR Data (Compound 4 in Acetone-ds)

| | | | ompound 4 | 4 in Acetone-d ₆) | | |
|----------|--------|-----------------|-----------|----------------------------------|--|--|
| Position | 13 | С | · | ¹H | | |
| 2 | 77.12 | СН | 5.59 | (1H, dd), J=2.7, 13.5 Hz | | |
| 3 | 43.49 | CH ₂ | 2.65 | (1H, dd), <i>J</i> =2.7, 17.2 Hz | | |
| | | | 3.14 | (1H, dd), J=13.5, 17.2 Hz | | |
| 4 | 197.65 | С | | | | |
| 5 | 162.27 | C | 12.47 | (-OH, s) | | |
| 6 | 108.96 | С | | | | |
| 7 . | 164.74 | C | | | | |
| 8 | 95.29 | СН | 6.04 | (1H, s) | | |
| 9 | 162.27 | C | | | | |
| 10 | 103.01 | C | | | | |
| 1' | 129.96 | С | | - | | |
| . 2' | 127.59 | С | | | | |
| 3, | 144.04 | С | | | | |
| 4' | 145.49 | С | | | | |
| 5' | 113.50 | СН | 6.82 | (1H. d), J=8.3 Hz | | |
| 6' | 118.51 | CH . | 6.96 | (1H, d), J=8.3 Hz | | |
| 1" | 25.13 | CH ₂ | 3.54 | (2H, d), J=6.5 Hz | | |
| 2" | 124.15 | СН | 5.18 | (1H, t), J=6.5 Hz | | |
| 3" | 135.34 | С | | | | |
| 4'' | 16.37 | CH ₃ | 1.64 | (3H, s) | | |
| . 5" | 40.36 | CH ₂ | 1.97 | (2H, t), J=7.0 Hz | | |
| 6'' | 27.35 | CH_2 | 2.04 | (2H, m) | | |
| . , 7" | 125.02 | СН | 5.06 | (1H, qt), J=1.2, 7.0 Hz | | |
| 8'' | 131.72 | С | | | | |
| 9" | 25.84 | CH ³ | 1.69 | (3H, s) | | |
| 10'' | 17.67 | CH ₃ | 1.55 | (3H;s) | | |
| 1" | 21.62 | CH ₂ | 3.26 | (2H, d), J=6.6 Hz | | |
| 2''' | 123.61 | СН | 5.24 | (1H, gt), $J=1.5$, 6.6 Hz | | |
| 3''' | 131.17 | С | | | | |
| 4''' | 25.75 | CH ₃ | 1.61 | (3H, s) | | |
| 5''' | 17.81 | CH ₃ | 1.76 | (3H, s) | | |

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